#### PATENT APPLICATION

#### TITLE

Use of N-octanoyl amino acids as slimming cosmetic and pharmaceutical active agent

#### APPLICANT

SOCIETE D'EXPLOITATION DE PRODUITS POUR LES INDUSTRIES CHIMIQUES SEPPIC

INVENTORS

Christine GARCIA

ABSTRACT

Use of a compound of formula (I):

$$R_1$$
-C(=O)-[NH-CH( $R_2$ )-C(=O)]<sub>m</sub>-OH (I)

in which  $R_1$  represents a hydrocarbon radical comprising 7 carbon atoms,  $R_2$  represents the characterizing chain of an amino acid and m is between 1 and 50, or of a mixture of the said compounds of formula (I), as a slimming active agent. Nontherapeutic method of treatment using the said compound and use of the said compound for preparing a medicament with lipolytic activity, intended to induce slimming of the human body.

The subject of the present invention is a novel use of cosmetic active agents for slimming the human body.

Some of the fat in the human body is stored in the form of triglycerides, in cells of the fatty tissue of the dermis, called adipocytes. Slimming reflects a reduction in the fat stored in the adipocytes. This process requires a preliminary step which takes place inside these cells and which consists in hydrolysing the triglycerides to fatty acids and glycerol. This phenomenon is called lipolysis.

Most slimming cosmetic formulations currently marketed contain at least one compound possessing a lipolytic activity. The one most frequently used is caffeine, but theophylline is also known to possess such a property.

During their search for novel active agents with lipolytic activity which have better compatibility with the skin than those of the state of the art, the inventors demonstrated that certain N-acylated derivatives of amino acids known for their soothing property also had a lipolytic property which was more effective than that of caffeine.

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Accordingly, according to a first aspect, the subject of the invention is the use of a compound of formula (I):

$$R_1$$
-C(=O)-[NH-CH( $R_2$ )-C(=O)]<sub>m</sub>-OH (!)

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in which  $R_1$  represents a linear or branched, saturated unsaturated, aliphatic hydrocarbon radical comprising 7 carbon atoms,  $R_2$ represents characterizing chain of an amino acid and m is between 1 and 50, or of a mixture of the said compounds of formula (I), as a slimming active agent, in а

composition containing a cosmetically acceptable medium.

The compound of formula (I) as defined above may be in the form of a free acid or in a partially or completely salified form. When the compound of formula (I) is in a salified form, it comprises in particular alkali metal salts such as the sodium, potassium or lithium salts, alkaline-earth metal salts such as the calcium, magnesium or strontium salts; an ammonium salt or a 10 salt of an amino alcohol such as the (2-hydroxyethyl)ammonium salt. It may also comprise metal salts such as divalent zinc or manganese salts, trivalent iron, lanthanum, cerium or aluminium salts. In general, the 15 degree of salification of the compound of formula (I) as defined above will additionally depend on its pKA and the salt concentration of the composition into which it is incorporated.

- In the following disclosure, the expression compound of formula (I) is understood to mean a compound of formula (I) in free form or in a partially or completely salified form.
- 25 The expression "characterizing chain" used to define the radical  $R_2$  denotes the nonfunctional principal chain of the amino acid considered.

Thus, for an amino acid represented by general formula 30 (IIIa):

$$H_2N-CH(R_2)-C(=O)-OH$$
 (IIIa)

and for a cyclic amino acid represented by formula 35 (IIIb):



the characterizing chain will be the chain represented by  $R_2$ .

R2 represents in particular the characterizing chain of an amino acid chosen from glycine, alanine, serine, valine, aspartic acid, glutamic acid, threonine, arginine, lysine, proline, leucine, phenylalanine, isoleucine, histidine, tyrosine, tryptophan, asparagine, glutamine, cysteine, cystine, methionine, hydroxyproline, hydroxylysine, sarcosine or ornithine.

The subject of the invention is mainly the use of a compound of formula (I) as defined above, in which, in the residue:

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$$-HN-CH(R_2)-C(=O)-$$
 (IIIa)

R<sub>2</sub> represents the characterizing chain of glycine.

- The subject of the invention is more particularly the use of a compound of formula (I) as defined above, in which m is a decimal number between 1 and 10 and it is preferably less than 5.
- 25 According to a most particular aspect of the present invention, in formula (I) as defined above, m is less than or equal to 2 and is more particularly less than or equal to 1.4.
- 30 According to another most particular aspect of the present invention, in formula (I) as defined above, m is equal to 1.

According to another particular variant of the present invention, a single compound of formula (I) as defined above is used in the composition containing the cosmetically acceptable medium.

According to another particular variant of the present invention, a mixture of compounds of formula (I) as defined above is used.

The compounds of formulae (I) are generally obtained by N-acylation of compounds of formulae (IIIa) or (IIIb), as defined above, or their salts.

In the case of a mixture of compounds of formulae (I)

10 it is for example obtained by N-acylation of the mixture of amino acids resulting from the total or partial hydrolysis of proteins of any origin.

These proteins may be of animal origin, such as for example collagen, elastin, fish flesh protein, fish gelatin, keratin or casein, of plant origin, such as cereal, flower or fruit proteins such as for example the proteins derived from soya bean, sunflower, oats, wheat, maize, barley, potato, lupin, field bean, sweet almond, kiwi, mango or apple; they may also be proteins obtained from Chorella (unicellular algae), pink algae, yeasts or silk.

This hydrolysis is carried out, for example, by heating to temperatures of between 60 and 130°C a protein placed in an acidic or alkaline medium.

This hydrolysis may also be carried out enzymatically, with a protease, optionally coupled with a post-30 alkaline or post-acid hydrolysis. When m is greater than 1, R<sub>2</sub> represents a single chain or several chains characterizing different amino acids, depending on the protein hydrolysed and the degree of hydrolysis.

35 The aminograms of a few proteins of plant origin are presented in the following tables:

Table 1

	Origin of the protein			
	(proportions of amino acids expressed			
	in wt %)			
	Oats	Soya	Wheat	Sunflower
		bean		
Glycine	6.9	4.2	3.2	6.2
Alanine	5.9	4.2	2.6	4.8
Serine	5.6	5.1	1.7	5.1
Aspartic acid	16.2	11.7	3.4	10.6
Glutamic acid	28.3	19.1	37.9	23.6
Valine	2.9	5.0	4.2	4.8
Threonine	3.1	3.9	2.7	4.4
Arginine	6.6	7.8	3.7	8.4
Lysine	3.6	6.2	1.9	3.2
Proline	4.7	5.4	11.7	3.0
Leucine	6.4	8.1	7.1	6.4
Phenylalanine	1.4	5.0	5.4	4.3
Isoleucine	2.2	4.8	3.7	4.1
Histidine	1.7	2.6	2.4	2.0
Tyrosine	1.5	3.5	3.1	2.7
Methionine	1.2	1.2	1.6	1.8
Cysteine/cystine	1.9	1.5	1.9	1.9
Tryptophan	_	1.0	1.0	1.3

Table 2

	Origin of the protein			
	(proportions of amino acids expressed			
		in wt %)		
	Lupin	Potato	Field	Maize
			bean	
Glycine	0.9	4.8	4.0	2.4
Alanine	2.4	5.0	4.0	7.95
Serine	6.1	5.8	4.9	5.1
Aspartic acid	15.8	12.5	10.5	10.6
Glutamic acid	8.0	11.5	16.8	23.6
Valine	7.9	7.1	4.5	4.8
Threonine	8.1	6.1	3.6	4.4

Arginine	16.1	5.0	9.21	8.4
Lysine	7.1	7.8	6.5	6.2
Proline	_	5.1	4.4	3.0
Leucine	7.45	10.4	7.4	8.1
Phenylalanine	8.6	6.4	4.4	4.3
Isoleucine	8.7	6.1	3.9	4.1
Histidine	-	2.2	2.6	2.0
Tyrosine	-	5.7	3.6	2.7
Methionine	0.6	2.4	0.8	1.8
Cysteine/cystine	_	1.6	1.7	1.9
Tryptophan	1.2	1.4	1.2	1.3
Ornithine	0.4	_	_	_

The acylation reaction is known to persons skilled in the art. It is described for example in international application published under the number WO 98/09611. It is carried out indifferently on an amino acid or on a mixture of amino acids. The acylating agent generally consists of an activated derivative of a carboxylic acid of formula  $R_1$ -C(=O)-OH, in which  $R_1$  is as defined above, such as a symmetric anhydride of this acid, the methyl ester of this acid, or an acid halide such as the acid chloride or the acid bromide.

The subject of the invention is also a nontherapeutic method of treating the human body intended for slimming it, characterized in that a composition containing a cosmetically acceptable medium and an effective quantity of at least one compound of formula (I) as defined above, is applied to it.

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- 20 The subject of the invention is also the use of at least one compound of formula (I), as defined above, for preparing a medicament with lipolytic activity, intended for inducing slimming of the human body.
- 25 In the compositions defined above, the compound of formula (I) is generally used in a quantity of between

0.01% and 10% of their weight, more particularly between 0.1% and 5% of their weight, and most particularly between 1% and 5% of their weight.

5 As the examples show, the compounds used in the cosmetic or therapeutic treatments defined above are characterized, unexpectedly, by a lipolytic activity greater than the compositions of the state of the art. They are therefore in general appropriate for the slimming treatments of the human body.

The compositions used in the said treatments are generally provided in the form of dilute aqueous or aqueous-alcoholic solutions, in the form of simple or multiple emulsions, such as water-in-oil (W/O), oil-in-water (O/W) or water-in-oil-in-water (W/O/W) emulsions in which the oil is of a vegetable or mineral nature, or in powdered form. They may also be dispersed or impregnated onto textile or onto nonwoven materials such as wipes, paper serviettes or clothing.

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The compositions used in the said treatments are administered to the subject in the conventional forms used in cosmetics and in pharmacy; this includes more particularly topical, oral or parenteral administrations.

In general, the compounds of formula (I) are combined with many types of adjuvants or active ingredients used 30 in cosmetic formulations, such as fatty substances, organic solvents, thickeners, gelling agents, antioxidants, emollients, opacifiers, stabilizers, foaming agents, perfumes, emulsifiers, which are ionic nonionic, fillers, sequestrants, chelators, preservatives, chemical screening agents or inorganic 35 screening agents, essential oils, colouring matter, pigments, hydrophilic or lipophilic active humectants, for example glycerin, preservatives, colorants, perfumes, cosmetic active agents, inorganic

or organic sunscreens, inorganic fillers such as iron oxides, titanium oxides and talc, synthetic fillers such as nylons and poly(methyl methacrylate) which are crosslinked or otherwise, silicone elastomers, sericites or plant extracts or alternatively lipid vesicles or any other ingredient customarily used in cosmetics.

As examples of oils which may be combined with the 10 compound of formula (I), there may be mentioned mineral oils such paraffin oil, liquid as paraffin, isoparaffins or white mineral oils, oils of animal origin, such as squalene or squalane, vegetable oils, such as sweet almond oil, copra oil, castor oil, jojoba 15 oil, olive oil, rapeseed oil, groundnut oil, sunflower oil, wheat germ oil, maize germ oil, soya bean oil, oil, lucerne oil, cottonseed seed poppy oil, pumpkinseed oil, evening primrose oil, millet oil, barley oil, rye oil, safflower oil, candlenut oil, 20 passionflower oil, hazelnut oil, palm oil, shea butter, apricot kernel oil, calophyllum oil, sysymbrium oil, avocado oil, calendula oil; ethoxylated vegetable oils; synthetic oils such as fatty acid esters such as butyl myristate, propyl myristate, cetyl myristate, isopropyl 25 palmitate, butyl stearate, hexadecyl stearate, isopropyl stearate, octyl stearate, isocetyl stearate, dodecvl oleate, hexyl laurate, propylene dicaprylate, esters derived from lanolic acid, such as isopropyl lanolate, isocetyl lanolate, monoglycerides, 30 diglycerides and triglycerides of fatty acids such as glyceryl triheptanoate, alkyl benzoates, poly-alphaolefins, polyolefins such as polyisobutene, synthetic isoalkanes such as isohexadecane, isododecane, fluorinated oils and silicone oils. Among the latter, there may be mentioned more particularly dimethylpoly-35 siloxanes, methylphenylpolysiloxanes, silicones modified with amines, silicones modified with fatty acids, silicones modified with alcohols, silicones modified with alcohols and fatty acids, silicones

modified with polyether groups, epoxy-modified silicones, silicones modified with fluorinated groups, cyclic silicones and silicones modified with alkyl groups.

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As other fatty substances which may be combined with this active agent, there may be mentioned fatty alcohols or fatty acids.

10 Among the thickening and/or emulsifying polymers used in the present invention are for example homopolymers copolymers of acrylic acid or of acrylic acid derivatives, homopolymers or copolymers of acrylamide, homopolymers or copolymers of acrylamide derivatives, 15 homopolymers or copolymers of acrylamidomethylpropaneacid, vinyl monomer, sulphonic trimethylaminoethylacrylate chloride, hydrocolloids of plant biosynthetic origin, for example xanthan gum, karaya gum, carrageenans, alginates; silicates; cellulose and 20 its derivatives; starch and its hydrophilic derivatives; polyurethanes. Among the polymers of the polyelectrolyte type which may be used in production of a gelled aqueous phase capable of being used in the preparation of W/O emulsions containing the 25 compounds of formula (I) which are the subject of the present invention, there are for example copolymers of acrylic acid and 2-methyl-[(1-oxo-2-propenyl)amino]-1propanesulphonic acid (AMPS), copolymers of acrylamide and 2-methyl-[(1-oxo-2-propenyl)amino]-1-propane-30 acid, sulphonic copolymers of 2-methyl-[(1-oxo-2propenyl)amino]-1-propanesulphonic acid and (2-hydroxyethyl) acrylate, homopolymer of 2-methyl-[(1oxo-2-propenyl)amino]-1-propanesulphonic homopolymer of acrylic acid, copolymers of acryloyl-35 ethyltrimethylammonium chloride and acrylamide, copolymers of AMPS and vinylpyrrolidone, copolymers of acrylic acid and alkyl acrylates whose carbon chain ten between and comprises thirty carbon atoms, copolymers of AMPS and alkyl acrylates whose carbon

chain comprises between ten and thirty carbon atoms. Such polymers are marketed respectively under the names SIMULGEL™ EG, SEPIGEL™ 305, SIMULGEL™ NS, SIMULGEL™ 800 and SIMULGEL™ A by the applicant.

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Among the waxes which can be used in the present invention, there may be mentioned for example beeswax; carnauba wax, candelilla wax; ouricoury wax; Japan wax; cork fibre or sugarcane wax; paraffin waxes; lignite waxes; microcrystalline waxes; lanolin wax; ozokerite; polyethylene wax; hydrogenated oils; silicone waxes; vegetable waxes; fatty alcohols and fatty acids which are solid at room temperature; glycerides which are solid at room temperature. Among the emulsifiers which can be used in the present invention, there may be mentioned for example fatty acids, ethoxylated fatty acids, fatty acid esters of sorbitol, ethoxylated fatty esters, polysorbates, polyglycerol ethoxylated fatty alcohols, sucrose esters, alkyl polyglycosides, sulphated and phosphated fatty alcohols or mixtures of alkyl polyglycosides and fatty alcohols described in French Patent Applications 2 668 080, 2 734 496, 2 756 195, 2 762 317, 2 784 680, 2 784 904, 2 791 565, 2 790 977, 2 807 435 and 2 804 432.

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As examples of an active ingredient which may be combined with the compound of formula (I) there may be mentioned compounds having a lightening or depigmenting action, such as for example arbutin, kojic ellagic acid, hydroquinone, vitamin C, magnesium ascorbyl phosphate, extracts of polyphenols, extracts, pine extracts, wine extracts, olive extracts, extracts, N-acylated proteins, N-acylated peptides, N-acylated amino acids, partial hydrolysates of N-acylated proteins, amino acids, peptides, total hydrolysates of proteins, partial hydrolysates proteins, polyols (for example glycerin or butylene glycol), urea, pyrrolidonecarboxylic acid derivatives of this acid, glycyrrhetinic acid, alpha-

bisabolol, sugars or sugar derivatives, polysaccharides or their derivatives, hydroxy acids, for example lactic acid, vitamins, vitamin derivatives such as retinol, vitamin E and its derivatives, minerals, enzymes, 5 coenzymes such as Coenzyme Q10, hormones or hormonesoya bean extracts, for substances, Raffermine™, wheat extracts, for example Tensine™ or Gliadine™, vegetable extracts such as extracts rich in tannins, extracts rich in isoflavones or extracts rich in terpenes, extracts of fresh water or marine algae, 10 essential waxes, bacterial extracts, minerals, lipids in general, lipids such as ceramides or phospholipids, active agents having a slimming action such as caffeine or its derivatives, active agents having antimicrobial activity or 15 purifying a action relation to greasy skins such as LIPACIDE™ PVB, active agents having an energizing or stimulating property such as SEPITONIC™ M3 or Physiogenyl™, panthenol and its derivatives such as SEPICAP™ MP, antiageing active 20 agents such as SEPILIFT<sup>TM</sup> DPHP, LIPACIDE<sup>TM</sup> SEPIVITAL™, moisturizing active agents SEPIVINOL™, such as SEPICALM™ S, SEPICALM™ VG and LIPACIDE™ DPHP, "antiphotoageing" antiageing active agents, agents protecting the integrity of the dermoepidermal 25 junction, active agents increasing the synthesis of the components of the extracellular matrix, active agents having a slimming, toning or draining activity such as caffeine, theophylline, cAMP, green tea, sage, ginko biloba, ivy, horse-chestnut, bamboo, ruscus, butcher's 30 broom, centella asiatica, heather, meadowsweet, fucus, rosemary, willow, active agents creating a sensation of skin, "heat" on the such as activators of microcirculation (for example nicotinates) or products creating a sensation of "freshness" on the skin (for example menthol and its derivatives). 35

As sunscreen which may be incorporated into the composition according to the invention, there may be

mentioned all those which appear in the amended cosmetics directive 76/768/EEC, annex VII.

The following experimental study illustrates the invention without, however, limiting it.

# A) Evaluation in vitro of the lipolytic activity of the compounds of formula (I)

### 10 (1) - Aim and principle of the method

The objective of the experiment is to demonstrate, in an in vitro model of isolated human adipocytes, the lipolytic activity of the compounds used. because triglycerides are stored in the adipocytes and constitute the fat. reserve. this For reserve diminish, which is the desired aim when slimming products are used, the triglycerides must be hydrolysed so that the fatty acids can be removed from the cell. Hydrolysis of the triglycerides to nonesterified fatty acids and glycerol is called lipolysis.

The method described consists in incubating the in the presence products of human adipocytes in suspension, followed by the measurement of the nonesterified acid level fatty in the adipocyte incubation medium. The free fatty acid level thus reflects the hydrolysis of the triglycerides present in the adipocytes.

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#### (2) - Experimental protocol

## (i) Cellular model:

35 The test is carried out using human adipocytes isolated and prepared as a cellular suspension. The adipocytes are isolated from the subcutaneous abdominal adipose tissue recovered during plastic surgery operations (abdominal plastic surgery operations) performed on

women. The cells are isolated from fresh tissue. The adipose tissue is isolated and dissociated by the action of a collagenase (SIGMATM, 1 mg/ml, 30 minutes at stirring). Collagenase 37°C, gentle digests connective tissue present in the adipose tissue. After digestion, the cells are filtered and washed in an appropriate culture medium containing MEM medium free of phenol red, free of glutamine (SIGMA) + 2.2 mg/ml of sodium bicarbonate (GIBCO) + 50 IU of penicillin (BIOWHITTAKER™) + 50  $\mu$ g/ml of streptomycin (BIOWHITTAKER™) 1왕 (v/v)of L-glutamine (BIOWHITTAKER™) + 0.5% of lipid-free serum albumin (SIGMA™). The adipocyte suspension is used immediately after its preparation.

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# (ii) Incubation of the products with the adipocytes

The test products are diluted in the adipocyte culture medium. They are incubated with the cells in suspension for two hours at 37°C (250  $\mu$ l of product + 250  $\mu$ l of adipocyte suspension).

#### (3) - Evaluation of the results

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After the incubation, the cell lysis is checked visually by the presence of a lipid layer surface of the cellular suspension. The supernatant media are collected. The free fatty acids are assayed by spectrophotometry using a commercial kit (NEFA $^{TM}$  C kit, WAKO), with reference to a fatty acid calibration series. The lypolytic activity of the products evaluated relative to a control group incubated in the presence of adipocytes and in the absence of product. The reactivity of the adipocytes is systematically checked for the measurement of the lipolytic activity of the reference products, caffeine (1,3,7-trimethylxanthine) and theophylline (1,3-dimethyl-2,6-dihydroxypurine). Five assays are performed for each of the test

products. The lipolytic activity of LIPACIDE<sup>TM</sup> C8G, which contains active material 100% N-octanoylglycine, was thus evaluated.

5 The results of the trials, expressed by the arithmetic means of the five assays carried out for each of the products, are presented in the following table:

	Incubation	Free fatty	Lipolytic
	concentration	acid	activity
	(wt %/volume)	concentration	(compared
		(μM)	with the
			control =
			100)
Control	_	11.54 ± 2.71	100
Caffeine	0.0019	15.10 ± 4.11	131
Theophylline	0.0019	16.53 ± 4.23	143
LIPACIDE™ C8G	0.0001	23.83 ± 5.11	206
LIPACIDE™ C8G	0.0005	25.30 ± 2.70	219
LIPACIDE™ C8G	0.0010	25.75 ± 0.86	223

10 These results show that while the slimming compositions of the state of the art (caffeine and theophylline) act on lipolysis with a multiplication factor of 1.3 to 1.4 relative to the control, that according to the invention, that is to say comprising at least one compound of formula (I) as defined above, acts with a factor of 2.06 to 2.23 at lower concentrations.

### B) - Examples of cosmetic formulations

20 In the following examples, the proportions are expressed as percentages by weight.

### Example 1: Slimming body milk

MONTANOV™ L	3.00%
Phytosqualane	8.00%
Sweet almond oil	2.00%

Water	qsp 100%
SEPIGEL™ 501	1.50%
LIPACIDE <sup>™</sup> C8G	3.00%
SEPICIDE™ CI	0.20%
SEPICIDE™ HB	0.30%
Perfume	0.30%
Example 2: Anti-sagging cream	(oval target
of the face)	
MONTANOV™ 202	3.50%
MONTANOV™ 14	1.00%
SEPILIFT <sup>TM</sup> DPHP	1.00%
LANOL™ 1688	15.00%
Wheat germ oil	5.00%
Water	qsp 100%
SIMULGEL™ EG	1.30%
LIPACIDE™ C8G	2.00%
SEPICIDE™ CI	0.20%
SEPICIDE™ HB	0.30%
Perfume	0.10%
Example 3: Anti-plumpness spra	ıy
MONTANE™ 60	3.30%
MONTANOX™ 60	1.70%
Caprylic/capric triglycerides	6.00%
Isohexadecane	5.00%
Magnesium Aluminium Silicate	1.50%
Water	qsp 100%
SIMULGEL™ EG	1.00%
LIPACIDE™ C8G	2.00%

Centalla asiatica/hydrocotyle 1.00% extract SEPICIDE<sup>TM</sup> CI 0.20% SEPICIDE<sup>TM</sup> HB 0.30% Perfume 0.40% Water qsp 100%

## Example 4: Refreshing slimming gel

SEPIGEL™ 305	3.50%
Hydroxyethylcellulose	1.00%
Caffeine	5.00%
Menthol	0.30%
Ethanol	50.00%
LIPACIDE™ C8G	3.00%
SEPICIDE™ LD	1.00%
Perfume	0.20%
Water	qsp 100%

## Example 5: Slimming body fluid

SIMULGEL™ NS	2.50%
Xanthan gum	0.20%
LANOL™ 99	5.00%
LIPACIDE™ C8G	2.00%
Ginkgo biloba extract	2.00%
Cola extract	1.00%
Ginseng extract	0.50%
SEPICIDE™ HB	1.50%
Perfume	0.10%
Water	qsp 100%

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# Example 6: Toning revitalizing lotion intended to be impregnated into body wipes

LIPACIDE™ C8G	1.50%
Glycerin	5.00%
Ethanol	5.00%
Ruscus extract	3.00%
SEPITONIC™ M3	1.00%

SEPICIDE™	CI	0.20	) ક
SEPICIDE™	НВ	0.30	) <b>%</b>
Water		asp	100%

### Example 7: Slimming shower gel

MONTALINE™ C40	8.00%
PROTEOL™ OAT	5.00%
Sodium lauryl sulphate	9.00%
LIPACIDE™ C8G	3.00%
Green tea extract	1.00%
KATHON™ CG	0.80
Green colorant	qs
Green tea perfume	1.00%
Lactic acid	qs pH=6.5
Water	qsp 100%

## Example 8: Biphasic disinfiltrating massage

Arabic coffee oil	1.00%
LANOL™ 189	20.00%
LANOL™ 99	10.00%
Borage oil	2.00%
Perfume	0.10%
LIPACIDE™ C8G	3.00%
Glycerin	3.00%
Ethanol	10.00%
Blue colorant	qs
Water	qsp 100%

The definitions of the commercial products used in the examples are the following:

SEPILIFT™ DPHP: (INCI name: Dipalmitoyl hydroxyproline), marketed by the company SEPPIC; SEPICIDE™ CI: Imidazoline urea (preservative), marketed by the company SEPPIC;

SEPICIDE™ HB: Mixture of phenoxyethanol, methylparaben, ethylparaben, propylparaben and butylparaben (preservative), marketed by the company SEPPIC;

SEPICIDE™ LD: Phenoxyethanol marketed by the company SEPPIC;

KATHON™ CG: (INCI name: methyl isothiazolinone / Methyl chloroisothiazolinone);

5 MONTANE™ 60: Sorbitan stearate;

MONTANOX™ 60: Polysorbate 60;

SIMULGEL $^{TM}$  EG: Self-reversible invert latex of copolymer such as those described in international publication WO 99/36445 (INCI name: Sodium acrylate/Sodium

acryloyldimethyl taurate copolymer and Isohexadecane and Polysorbate 80) marketed by the company SEPPIC; SIMULGEL™ NS: Self-reversible invert latex of copolymer such as those described in international publication WO 99/36445 (INCI name: hydroxyethylacrylate/Sodium

15 acryloyldimethyl taurate copolymer and squalane and Polysorbate 60) marketed by the company SEPPIC;

SEPIGEL™ 305: Self-reversible invert latex (INCI name:

SEPIGEL™ 305: Self-reversible invert latex (INCI name: Polyacrylamide / C13-14 Isoparaffin / Laureth-7);

SEPIGEL™ 501: Self-reversible invert latex INCI name:

20 C13-14 Isoparaffin/Mineral Oil/Sodium polyacrylate-/Polyacrylamide/Polysorbate 85;

LANOL™ 99: Isononyl isononanoate marketed by the company SEPPIC;

LANOL™ 189: Isostearyl isononanoate

25 LANOL™ 1688: Cetearyl ethyl hexanoate marketed by the company SEPPIC;

SEPITONIC™ M3: Mixture of magnesium aspartate, copper gluconate and zinc gluconate marketed by the company SEPPIC;

30 MONTALINE™ C40: Cocamidopropyl betainamide MEA chloride PROTEOL™ OAT: Sodium Lauroyl oat amino acids;

MONTANOV™ 14: Myristyl alcohol / Myristyl glucoside;

 ${\tt MONTANOV^{TM}}$  L: Emulsifying agent based on a C14-C22 alcohol and a C12-C20 alkyl polyglucoside such as those

described in European Patent Application EP 0 995 487;

MONTANOV™ 202 is an emulsifying agent based on arachidyl alcohol, behenyl alcohol and arachidyl polyglucoside.